

# Supporting Information

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Table S1. Nucleotide sequences of primers used for site-directed mutagenesis

Mutants	Primer sequences (5'-3')
Y252F	CCCTGCCCCACCT <b>T</b> CAACAACCACCT <b>G</b> TACAAACAAATTTCCAGCC Tyr→Phe BsrGI
Y272F	CCAATCAGGAG <b>GCT</b> TCGAACGACAATCACT <b>T</b> CTTTGGCTACAG BstBI Tyr→Phe
Y444F	CGACCAGTACCTGTATT <b>CTTAAG</b> CAGAACAAACACTCCAAG Tyr→Phe AflII
Y500F	CAACAACAGTGAAT <b>TCT</b> CGTGG <b>ACCGGT</b> GCTACCAAGTACC Tyr→Phe AgeI
Y700F	GGAATCCCGAAATTCAGT <b>TCACTTCGAACT</b> TACAACAAGTCTG Tyr→Phe BstBI
Y704F	GGAATCCCGAAATTCAGTACACT <b>TTCGAACTTCAACAAGTCTG</b> BstBI Tyr→Phe
Y730F	CCTCGCCCCATT <b>GGT</b> ACCAGAT <b>T</b> CCTGACTCGTAATC Acc65I Tyr→Phe

Nucleotide sequences of PCR primers used for site-directed mutagenesis of surface-exposed tyrosine residues. The codon triplets are shown in bold; red denotes that the mutations from tyrosine to phenylalanine residues, and green indicates that silent mutations to create the restriction enzyme sites shown (underlined), which were used to obtain the desired clones.

**Table S2. Titers of the WT and Y-F mutant scAAV2 vectors**

AAV vectors	Packaging titers, vgs/ml			
	First	Second	Third	Fourth
WT scAAV2-EGFP	$3.4 \times 10^{11}$	$1.0 \times 10^{12}$	$3.2 \times 10^{11}$	$3.0 \times 10^{11}$
Y252F scAAV2-EGFP	$3.8 \times 10^{11}$	$4.0 \times 10^{11}$	ND	ND
Y272F scAAV2-EGFP	$7.7 \times 10^{11}$	$1.0 \times 10^{11}$	ND	ND
Y444F scAAV2-EGFP	$9.7 \times 10^{10}$	$4.0 \times 10^{10}$	$6.0 \times 10^9$	$5.0 \times 10^{10}$
Y500F scAAV2-EGFP	$8.8 \times 10^{10}$	$2.0 \times 10^9$	$4.0 \times 10^{10}$	$6.0 \times 10^{10}$
Y700F scAAV2-EGFP	$1.0 \times 10^{11}$	$4.0 \times 10^{11}$	ND	ND
Y704F scAAV2-EGFP	$6.0 \times 10^{11}$	$2.0 \times 10^{11}$	ND	ND
Y730F scAAV2-EGFP	$1.2 \times 10^{11}$	$5.0 \times 10^{11}$	$1.2 \times 10^{11}$	$4.0 \times 10^{11}$

ND, not done. Physical particle titers of the WT and surface-exposed tyrosine-mutant scAAV2-EGFP vectors generated by four separate packaging runs. The first two packaging runs were performed by using one cell factory for each of the vectors, and the last two packaging runs were carried out by using five 150-mm culture dishes for each vector. Vector titers were determined on DNA slot-blots using a  $^{32}\text{P}$ -labeled EGFP DNA as a probe.